50C

STIC Biotechnology Systems Branch

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Application Serial Number: 10/516,490A
Source: PCT/10
Date Processed by STIC: 1/16/06

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1. EFS-Bio (<**http://www.uspto.gov/ebc/efs/downloads/documents.htm**> , **EFS Submission User Manual** - ePAVE)
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Revised 01/24/05

PCT

RAW SEQUENCE LISTING DATE: 01/10/2006
PATENT APPLICATION: **US/10/516,490A** TIME: 08:39:08

Input Set : **A:\X15642.NatlPhase.ST25.txt**
Output Set: **N:\CRF4\01102006\J516490A.raw**

```
      3 <110> APPLICANT: Richard Dennis DiMarchi
      4       David Lee Smiley
      5       Lianshan Zhang
      7 <120> TITLE OF INVENTION: MODIFIED GLUCAGON-LIKE PEPTIDE-1 ANALOGS
      9 <130> FILE REFERENCE: X-15642 National Phase
C--> 11 <140> CURRENT APPLICATION NUMBER: US/10/516,490A
C--> 11 <141> CURRENT FILING DATE: 2004-12-01
     11 <160> NUMBER OF SEQ ID NOS: 24
     13 <170> SOFTWARE: PatentIn version 3.2
     15 <210> SEQ ID NO: 1
     16 <211> LENGTH: 31
     17 <212> TYPE: PRT
     18 <213> ORGANISM: Artificial
     20 <220> FEATURE:
     21 <223> OTHER INFORMATION: Synthetic constructs
     24 <220> FEATURE:
     25 <221> NAME/KEY: MISC_FEATURE
     26 <222> LOCATION: (1)..(1)
     27 <223> OTHER INFORMATION: Xaa= L-histidine, D-histidine, desamino-histidine,
     28       2-amino-histidine, beta-hydroxy-
     29       histidine, homohistidine, alpha-fluoromethyl-histidine, or alpha
     30       methyl-histidine
     32 <220> FEATURE:
     33 <221> NAME/KEY: MISC_FEATURE
     34 <222> LOCATION: (2)..(2)
     35 <223> OTHER INFORMATION: Xaa= Ala, Gly, Val, Leu, Ile, Ser, or Thr
     37 <220> FEATURE:
     38 <221> NAME/KEY: MISC_FEATURE
     39 <222> LOCATION: (6)..(6)
     40 <223> OTHER INFORMATION: Xaa= Phe, Trp, or Tyr
     42 <220> FEATURE:
     43 <221> NAME/KEY: MISC_FEATURE
     44 <222> LOCATION: (10)..(10)
     45 <223> OTHER INFORMATION: Xaa= Val, Trp, Ile, Leu, Phe, or Tyr
     47 <220> FEATURE:
     48 <221> NAME/KEY: MISC_FEATURE
     49 <222> LOCATION: (12)..(12)
     50 <223> OTHER INFORMATION: Xaa= Ser, Trp, Tyr, Phe, Lys, Ile, Leu, Val
     52 <220> FEATURE:
     53 <221> NAME/KEY: MISC_FEATURE
     54 <222> LOCATION: (13)..(13)
     55 <223> OTHER INFORMATION: Xaa= Tyr, Trp, or Phe
     57 <220> FEATURE:
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pr 2-3, 5-7

Does Not Comply
Corrected Diskette Needed

RAW SEQUENCE LISTING
PATENT APPLICATION: US/10/516,490A

DATE: 01/10/2006
TIME: 08:39:08

Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

58 <221> NAME/KEY: MISC_FEATURE
59 <222> LOCATION: (14)..(14)
60 <223> OTHER INFORMATION: Xaa= Leu, Phe, Tyr, or Trp
62 <220> FEATURE:
63 <221> NAME/KEY: MISC_FEATURE
64 <222> LOCATION: (16)..(16)
65 <223> OTHER INFORMATION: Xaa= Gly, Glu, Asp, Lys
67 <220> FEATURE:
68 <221> NAME/KEY: MISC_FEATURE
69 <222> LOCATION: (19)..(19)
70 <223> OTHER INFORMATION: Xaa= Ala, Val, Ile, or Leu
72 <220> FEATURE:
73 <221> NAME/KEY: MISC_FEATURE
74 <222> LOCATION: (21)..(21)
75 <223> OTHER INFORMATION: Xaa= Glu, Ile, or Ala
77 <220> FEATURE:
78 <221> NAME/KEY: MISC_FEATURE
79 <222> LOCATION: (24)..(24)
80 <223> OTHER INFORMATION: Xaa= Ala, or Glu
82 <220> FEATURE:
83 <221> NAME/KEY: MISC_FEATURE
84 <222> LOCATION: (27)..(27)
85 <223> OTHER INFORMATION: Xaa= Val, or Ile
87 <220> FEATURE:
88 <221> NAME/KEY: MISC_FEATURE
89 <222> LOCATION: (31)..(31)
90 <223> OTHER INFORMATION: Xaa= L-Cys, D-Cys, homocysteine, or penicillamine
92 <400> SEQUENCE: 1

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W--> 94 Xaa Xaa Glu Gly Thr Xaa Thr Ser Asp Xaa Ser Xaa Xaa Xaa Glu Xaa
     95 1               5                  10                  15
W--> 98 Gln Ala Xaa Lys Xaa Phe Ile Xaa Trp Leu Xaa Lys Gly Arg Xaa
     99             20                  25                  30
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102 <210> SEQ ID NO: 2
103 <211> LENGTH: 31
104 <212> TYPE: PRT
105 <213> ORGANISM: Artificial
107 <220> FEATURE:
108 <223> OTHER INFORMATION: Synthetic construct
111 <220> FEATURE:
112 <221> NAME/KEY: MISC_FEATURE
113 <222> LOCATION: (1)..(1)
114 <223> OTHER INFORMATION: Xaa= L-histidine, D-histidine, desamino-histidine,
115 2-amino-histidien, beta-hydroxy-
116 histidine, homohistidine, alpha-fluoromethyl-histidine, or
117 alpha-methyl-histidine
119 <220> FEATURE:
120 <221> NAME/KEY: MISC_FEATURE
121 <222> LOCATION: (2)..(2)
122 <223> OTHER INFORMATION: Xaa= Gly, Ala, Val, Leu, Ile, Ser or Thr

RAW SEQUENCE LISTING DATE: 01/10/2006
PATENT APPLICATION: US/10/516,490A TIME: 08:39:08

Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

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     124 <220> FEATURE:
     125 <221> NAME/KEY: MISC_FEATURE
     126 <222> LOCATION: (10)..(10)
     127 <223> OTHER INFORMATION: Xaa = Val, Phe, Tyr, or Trp
     129 <220> FEATURE:
     130 <221> NAME/KEY: MISC_FEATURE
     131 <222> LOCATION: (12)..(12)
     132 <223> OTHER INFORMATION: Xaa = Ser, Tyr, Trp, Phe, Lys, Ile, Leu, or Val
     134 <220> FEATURE:
     135 <221> NAME/KEY: MISC_FEATURE
     136 <222> LOCATION: (16)..(16)
     137 <223> OTHER INFORMATION: Xaa = Gly, Clu, Asp, or Lys
     139 <220> FEATURE:
     140 <221> NAME/KEY: MISC_FEATURE
     141 <222> LOCATION: (19)..(19)
     142 <223> OTHER INFORMATION: Xaa = Ala, Val, Ile, or Leu
     144 <220> FEATURE:
     145 <221> NAME/KEY: MISC_FEATURE
     146 <222> LOCATION: (27)..(27)
     147 <223> OTHER INFORMATION: Xaa = Val or Ile
     149 <220> FEATURE:
     150 <221> NAME/KEY: MISC_FEATURE
     151 <222> LOCATION: (31)..(31)
     152 <223> OTHER INFORMATION: Xaa = L-Cys, D-Cys, homocysteine, or
penicillamine
     154 <400> SEQUENCE: 2
W--> 156 Xaa Xaa Glu Gly Thr Phe Thr Ser Asp Xaa Ser Xaa Tyr Leu Glu Xaa
     157 1               5                   10                  15
W--> 160 Gln Ala Xaa Lys Glu Phe Ile Ala Trp Leu Xaa Lys Gly Arg Xaa
     161             20                  25                  30
     164 <210> SEQ ID NO: 3
     165 <211> LENGTH: 42
     166 <212> TYPE: PRT
     167 <213> ORGANISM: Artificial
     169 <220> FEATURE:
     170 <223> OTHER INFORMATION: Synthetic construct
     173 <220> FEATURE:
     174 <221> NAME/KEY: MISC_FEATURE
     175 <222> LOCATION: (1)..(1)
     176 <223> OTHER INFORMATION: Xaa = L-histidine, D-histidine, desamino-
histidine,
     177       2-amino-histidine, beta-hydroxy-
     178       histidine, homohistidine, alpha-fluoromethyl-histidine, or
     179       alpha-methyl-histidine
     181 <220> FEATURE:
     182 <221> NAME/KEY: MISC_FEATURE
     183 <222> LOCATION: (2)..(2)
     184 <223> OTHER INFORMATION: Xaa = Ala, Gly, Val, Leu, Ile, Ser, or Thr
     186 <220> FEATURE:
     187 <221> NAME/KEY: MISC_FEATURE
     188 <222> LOCATION: (6)..(6)
```

do you mean Glu?

RAW SEQUENCE LISTING DATE: 01/10/2006
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Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

189 <223> OTHER INFORMATION: Xaa = Phe, Trp, or Tyr
191 <220> FEATURE:
192 <221> NAME/KEY: MISC_FEATURE
193 <222> LOCATION: (10)..(10)
194 <223> OTHER INFORMATION: Xaa = Val, Trp, Ile, Leu, Phe, or Tyr
196 <220> FEATURE:
197 <221> NAME/KEY: MISC_FEATURE
198 <222> LOCATION: (12)..(12)
199 <223> OTHER INFORMATION: Xaa = Ser, Trp, Tyr, Phe, Lys, Ile, Leu, Val
201 <220> FEATURE:
202 <221> NAME/KEY: MISC_FEATURE
203 <222> LOCATION: (13)..(13)
204 <223> OTHER INFORMATION: Xaa = Tyr, Trp, or Phe
206 <220> FEATURE:
207 <221> NAME/KEY: MISC_FEATURE
208 <222> LOCATION: (14)..(14)
209 <223> OTHER INFORMATION: Xaa = Leu, Phe, Tyr, or Trp
211 <220> FEATURE:
212 <221> NAME/KEY: MISC_FEATURE
213 <222> LOCATION: (16)..(16)
214 <223> OTHER INFORMATION: Xaa = Gly, Glu, Asp, or Lys
216 <220> FEATURE:
217 <221> NAME/KEY: MISC_FEATURE
218 <222> LOCATION: (19)..(19)
219 <223> OTHER INFORMATION: Xaa = Ala, Val, Ile, or Leu
221 <220> FEATURE:
222 <221> NAME/KEY: MISC_FEATURE
223 <222> LOCATION: (21)..(21)
224 <223> OTHER INFORMATION: Xaa = Glu, Ile, or Ala
226 <220> FEATURE:
227 <221> NAME/KEY: MISC_FEATURE
228 <222> LOCATION: (24)..(24)
229 <223> OTHER INFORMATION: Xaa = Ala or Glu
231 <220> FEATURE:
232 <221> NAME/KEY: MISC_FEATURE
233 <222> LOCATION: (27)..(27)
234 <223> OTHER INFORMATION: Xaa = Val or Ile
236 <220> FEATURE:
237 <221> NAME/KEY: MISC_FEATURE
238 <222> LOCATION: (28)..(28)
239 <223> OTHER INFORMATION: Xaa = Lys, Asp, Arg, or Glu
241 <220> FEATURE:
242 <221> NAME/KEY: MISC_FEATURE
243 <222> LOCATION: (30)..(30)
244 <223> OTHER INFORMATION: Xaa = Gly, Pro, or Arg
246 <220> FEATURE:
247 <221> NAME/KEY: MISC_FEATURE
248 <222> LOCATION: (31)..(31)
249 <223> OTHER INFORMATION: Xaa = Gly, Pro, Ser, L-Cys, D-Cys, homocysteine, or penicillamine

RAW SEQUENCE LISTING DATE: 01/10/2006
PATENT APPLICATION: US/10/516,490A TIME: 08:39:08

Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

```
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     252 <221> NAME/KEY: MISC_FEATURE
     253 <222> LOCATION: (32)..(32)
     254 <223> OTHER INFORMATION: Xaa = Ser, Pro, His, L-Cys, D-Cys, homocysteine,
penicillamine,
     255       NH2
     257 <220> FEATURE:
     258 <221> NAME/KEY: MISC_FEATURE
     259 <222> LOCATION: (33)..(33)
     260 <223> OTHER INFORMATION: Xaa = Ser, Arg, Thr, Trp, Lys, L-Cys, D-Cys,
homocysteine,
     261       penicillamine, NH2 or
     262       is absent
     264 <220> FEATURE:
     265 <221> NAME/KEY: MISC_FEATURE
     266 <222> LOCATION: (34)..(34)
     267 <223> OTHER INFORMATION: Xaa = Ser, Gly, L-Cys, D-Cys, homocysteine,
penicillamine, NH2,
     268       or is absent
     270 <220> FEATURE:
     271 <221> NAME/KEY: MISC_FEATURE
     272 <222> LOCATION: (35)..(35)
     273 <223> OTHER INFORMATION: Xaa = Ala, Asp, Arg, Glu, Lys, Gly, L-Cys, D-Cys,
homocysteine,
     274       penicillamine,
     275       NH2 or is absent
     277 <220> FEATURE:
     278 <221> NAME/KEY: MISC_FEATURE
     279 <222> LOCATION: (36)..(36)
     280 <223> OTHER INFORMATION: Xaa = Pro, Ala, L-Cys, D-Cys, homocysteine,
penicillamine, NH2,
     281       or is absent
     283 <220> FEATURE:
     284 <221> NAME/KEY: MISC_FEATURE
     285 <222> LOCATION: (37)..(37)
     286 <223> OTHER INFORMATION: Xaa = Pro, Ala, L-Cys, D-Cys, homocysteine,
penicillamine, NH2 or
     287       is absent
     289 <220> FEATURE:
     290 <221> NAME/KEY: MISC_FEATURE
     291 <222> LOCATION: (38)..(38)
     292 <223> OTHER INFORMATION: Xaa = Pro, Ala, Arg, Lys, His, L-Cys, D-Dys,
homocysteine,
     293       penicillamine, NH2 or
     294       is absent
     296 <220> FEATURE:
     297 <221> NAME/KEY: MISC_FEATURE
     298 <222> LOCATION: (39)..(39)
     299 <223> OTHER INFORMATION: Xaa = Ser, His, Pro, Lys, Arg, L-Cys, D-Cys,
homocysteine,
     300       penicillamine, NH2 or
     301       is absent
     303 <220> FEATURE:
```

do you mean "amidation"? If so, please add "amidated" to "NH_2"

Cys?

304 <221> NAME/KEY: MISC_FEATURE
305 <222> LOCATION: (40)..(40)
306 <223> OTHER INFORMATION: Xaa = His, Ser, Arg, Lys, L-Cys, D-Cys, homocysteine,
307 penicillamine, NH2 or

Please ensure that amino acids are spelled correctly in subsequent sequences.

RAW SEQUENCE LISTING ERROR SUMMARY DATE: 01/10/2006
PATENT APPLICATION: US/10/516,490A TIME: 08:39:09

Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

FYI

Please Note:
Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220>
to <223> fields of each sequence which presents at least one n or Xaa.

Seq#:1; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,31
Seq#:2; Xaa Pos. 1,2,10,12,16,19,27,31
Seq#:3; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,28,30,31,32,33,34,35,36
Seq#:3; Xaa Pos. 37,38,39,40,41,42
Seq#:4; Xaa Pos. 1,2,10,16,19,27,28,30,31,32,33,34,35,36,37,38,39,40,41,42
Seq#:5; Xaa Pos. 1,2,16,19,27,32,33,34,35,36,37,38,39,40,41,42
Seq#:6; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,28,30,31,32,33,34,35,36
Seq#:6; Xaa Pos. 37,38,39,40,41,42,43,44,45
Seq#:7; Xaa Pos. 32,33,34,35,36,37,38,39,40,41,42,43,44,45
Seq#:8; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27
Seq#:9; Xaa Pos. 1,2,10,12,16,19,27
Seq#:10; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,28,30,31,32,33,34,35,36
Seq#:10; Xaa Pos. 37,38,39,40,41,42
Seq#:11; Xaa Pos. 1,2,10,16,19,27,28,30,31,32,33,34,35,36,37,38,39,40,41,42
Seq#:12; Xaa Pos. 1,2,16,19,27,32,33,34,35,36,37,38,39,40,41,42
Seq#:13; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,28,30,31,32,33,34,35,36
Seq#:13; Xaa Pos. 37,38,39,40,41,42,43,44,45
Seq#:14; Xaa Pos. 32,33,34,35,36,37,38,39,40,41,42,43,44,45
Seq#:15; Xaa Pos. 1,2,6,10,12,13,14,16,19,21,24,27,31

Invalid <213> Response:
Use of "Artificial" only as "<213> Organism" response is incomplete, per 1.823(b) of New Sequence Rules. Valid response is Artificial Sequence.

Seq#:1,2,3,4,5,6,7,8,9,10,11,12,13,14,15,16,17,18,19,20,21,22

VERIFICATION SUMMARY DATE: 01/10/2006
PATENT APPLICATION: US/10/516,490A TIME: 08:39:09

Input Set : A:\X15642.NatlPhase.ST25.txt
Output Set: N:\CRF4\01102006\J516490A.raw

L:11 M:270 C: Current Application Number differs, Replaced Current Application No
L:11 M:271 C: Current Filing Date differs, Replaced Current Filing Date
L:94 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1 after pos.:0
L:98 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:1 after pos.:16
L:156 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:0
L:160 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:16
L:324 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 after pos.:0
L:328 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 after pos.:16
L:332 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 after pos.:32
L:466 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4 after pos.:0
L:470 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4 after pos.:16
L:474 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:4 after pos.:32
L:583 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:5 after pos.:0
L:587 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:5 after pos.:16
L:591 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:5 after pos.:32
L:762 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:6 after pos.:0
L:766 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:6 after pos.:16
L:770 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:6 after pos.:32
L:869 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:7 after pos.:16
L:873 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:7 after pos.:32
L:951 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:8 after pos.:0
L:955 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:8 after pos.:16
L:1008 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:9 after pos.:0
L:1012 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:9 after pos.:16
L:1160 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10 after pos.:0
L:1164 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10 after pos.:16
L:1168 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:10 after pos.:32
L:1286 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:11 after pos.:0
L:1290 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:11 after pos.:16
L:1294 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:11 after pos.:32
L:1391 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:12 after pos.:0
L:1395 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:12 after pos.:16
L:1399 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:12 after pos.:32
L:1561 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:13 after pos.:0
L:1565 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:13 after pos.:16
L:1569 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:13 after pos.:32
L:1658 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:14 after pos.:16
L:1662 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:14 after pos.:32
L:1745 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:15 after pos.:0
L:1749 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:15 after pos.:16